

SEQUENCE LISTING

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Mutated SRC Oncogene Composition and Methods <120>

USF-T136 <130>

<140> US 09/444,711 1999-11-24 <141>

<160>

<170> PatentIn version 3.1

<210>

<211> 1611

<212> DNA

<213> Homo sapiens

<220>

misc_feature

<222> (1)..(1611)

nucleotide sequence of normal c-Src oncogene coding region <223>

<220>

<221> CDS

<222> (1)..(1611)

<223>

<400> 1

atg ggt agc aac aag agc aag ccc aag gat gcc agc cag cgg cgc cgc 48 Met Gly Ser Asn Lys Ser Lys Pro Lys Asp Ala Ser Gln Arg Arg Arg

age etg gag eee gee gag aac gtg eae gge get gge gge get tte 96 Ser Leu Glu Pro Ala Glu Asn Val His Gly Ala Gly Gly Ala Phe

ccc gcc tcg cag acc ccc agc aag cca gcc tcg gcc gac ggc cac cgc Pro Ala Ser Gln Thr Pro Ser Lys Pro Ala Ser Ala Asp Gly His Arg

ggc ccc agc gcg gcc ttc gcc ccc gcg gcc gcc gag ccc aag ctg ttc Gly Pro Ser Ala Ala Phe Ala Pro Ala Ala Ala Glu Pro Lys Leu Phe

50 55

40

192

144

RECEIVED

NOV 1 9 2002

					tcg Ser 70							_		240
					gtg Val									288
		_			gac Asp	_		-					_	336
					gag Glu									384
					tac Tyr									432
					gag Glu 150									480
					ctc Leu									528
	_	_	_		acc Thr	_								576
					aag Lys									624
					ggc Gly									672
_	Leu	_	_	_	gtg Val 230	_				_	_		_	720
					acc Thr									768
					gcc Ala									816

														gly ggg		864
														ggc Gly		912
_				_		_	_		_	_	-	_	_	aag Lys	-	960
														gag Glu 335		1008
			_	_			_	_						gac Asp		1056
														gtg Val		1104
														atg Met		1152
	-			-										gag Glu		1200
														gaa Glu 415		1248
			_											tgg Trp		1296
_		_	_	_				_				_	_	gac Asp		1344
					_	_						_		cgg Arg		1392
														gag Glu		1440
			_		_	_	_		_				_	cac His 495	_	1488

4

ctc atg tgc cag tgc tgg cgg aag gag cct gag gag cgg ccc acc ttc

Leu Met Cys Gln Cys Trp Arg Lys Glu Pro Glu Glu Arg Pro Thr Phe
500

gag tac ctg cag gcc ttc ctg gag gac tac ttc acg tcc acc gag ccc
Glu Tyr Leu Gln Ala Phe Leu Glu Asp Tyr Phe Thr Ser Thr Glu Pro
515

cag tac cag ccc ggg gag aac ctc tag

Cag tac cag ccc ggg gag aac ctc tag

Gln Tyr Gln Pro Gly Glu Asn Leu
530

1536

1536

1536

1536

1536

1536

1536

1537

1611

<210> 2

<211> 536

<212> PRT

<213> Homo sapiens

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<221> MISC_FEATURE

<222> (1)..(536)

<223> amino acid sequence of non-receptor tyrosine kinase encoded
 by the normal c-Src coding region

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Met Gly Ser Asn Lys Ser Lys Pro Lys Asp Ala Ser Gln Arg Arg 1 5 10 15

Ser Leu Glu Pro Ala Glu Asn Val His Gly Ala Gly Gly Ala Phe 20 25 30

Pro Ala Ser Gln Thr Pro Ser Lys Pro Ala Ser Ala Asp Gly His Arg
35 40 45

Gly Pro Ser Ala Ala Phe Ala Pro Ala Ala Ala Glu Pro Lys Leu Phe 50 55 60

Gly Gly Phe Asn Ser Ser Asp Thr Val Thr Ser Pro Gln Arg Ala Gly 65 70 75 80

Pro Leu Ala Gly Gly Val Thr Thr Phe Val Ala Leu Tyr Asp Tyr Glu 85 90 95

Ser Arg Thr Glu Thr Asp Leu Ser Phe Lys Lys Gly Glu Arg Leu Gln
100 105 110

Ile Val Asn Asn Thr Glu Gly Asp Trp Trp Leu Ala His Ser Leu Ser
115 120 125

Thr Gly Gln Thr Gly Tyr Ile Pro Ser Asn Tyr Val Ala Pro Ser Asp 130 135 140

Ser Ile Gln Ala Glu Glu Trp Tyr Phe Gly Lys Ile Thr Arg Arg Glu 145 150 155 160

CM

Ser Glu Arg Leu Leu Asn Ala Glu Asn Pro Arg Gly Thr Phe Leu 165 170 175

Val Arg Glu Ser Glu Thr Thr Lys Gly Ala Tyr Cys Leu Ser Val Ser 180 185 190

Asp Phe Asp Asn Ala Lys Gly Leu Asn Val Lys His Tyr Lys Ile Arg 195 200 205

Lys Leu Asp Ser Gly Gly Phe Tyr Ile Thr Ser Arg Thr Gln Phe Asn 210 215 220

Ser Leu Gln Gln Leu Val Ala Tyr Tyr Ser Lys His Ala Asp Gly Leu 225 230 235 240

Cys His Arg Leu Thr Thr Val Cys Pro Thr Ser Lys Pro Gln Thr Gln 245 250 255

Gly Leu Ala Lys Asp Ala Trp Glu Ile Pro Arg Glu Ser Leu Arg Leu 260 265 270

Glu Val Lys Leu Gly Gln Gly Cys Phe Gly Glu Val Trp Met Gly Thr 275 280 285

Trp Asn Gly Thr Thr Arg Val Ala Ile Lys Thr Leu Lys Pro Gly Thr 290 295 300

Met Ser Pro Glu Ala Phe Leu Gln Glu Ala Gln Val Met Lys Lys Leu 305 310 315 320

Arg His Glu Lys Leu Val Gln Leu Tyr Ala Val Val Ser Glu Glu Pro 325 330 335

Ile Tyr Ile Val Thr Glu Tyr Met Ser Lys Gly Ser Leu Leu Asp Phe

Leu Lys Gly Glu Thr Gly Lys Tyr Leu Arg Leu Pro Gln Leu Val Asp 355 360 365

Met Ala Ala Gln Ile Ala Ser Gly Met Ala Tyr Val Glu Arg Met Asn 370 375 380

Tyr Val His Arg Asp Leu Arg Ala Ala Asn Ile Leu Val Gly Glu Asn 385 390 395 400

Leu Val Cys Lys Val Ala Asp Phe Gly Leu Ala Arg Leu Ile Glu Asp 405 410 415

Asn Glu Tyr Thr Ala Arg Gln Gly Ala Lys Phe Pro Ile Lys Trp Thr 420 425 430

Ala Pro Glu Ala Ala Leu Tyr Gly Arg Phe Thr Ile Lys Ser Asp Val 435 440 445

Trp Ser Phe Gly Ile Leu Leu Thr Glu Leu Thr Thr Lys Gly Arg Val 450 455 460



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Pro Tyr Pro Gly Met Val Asn Arg Glu Val Leu Asp Gln Val Glu Arg
Gly Tyr Arg Met Pro Cys Pro Pro Glu Cys Pro Glu Ser Leu His Asp
                                    490
Leu Met Cys Gln Cys Trp Arg Lys Glu Pro Glu Glu Arg Pro Thr Phe
            500
                                505
Glu Tyr Leu Gln Ala Phe Leu Glu Asp Tyr Phe Thr Ser Thr Glu Pro
                            520
Gln Tyr Gln Pro Gly Glu Asn Leu
<210> 3
<211> 1611
<212> DNA
<213> Homo sapiens
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<223> nucleotide sequence of mutant c-Src oncogene coding region
<220>
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      CDS
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       (1)..(1593)
<223>
<220>
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      misc feature
<222>
      (1591)..(1591)
<223> Point mutation in normal c-Src causes transition from c-->t and
       the formation of a stop codon.
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atg ggt agc aac aag agc aag ccc aag gat gcc agc cag cgg cgc cgc
Met Gly Ser Asn Lys Ser Lys Pro Lys Asp Ala Ser Gln Arg Arg Arg
1
                5
                                                        15
age etg gag eee gee gag aac gtg cae gge get gge gge get tte
                                                                       96
Ser Leu Glu Pro Ala Glu Asn Val His Gly Ala Gly Gly Ala Phe
            20
                                                    30
ccc gcc tcg cag acc ccc agc aag cca gcc tcg gcc gac ggc cac cgc
                                                                     144
Pro Ala Ser Gln Thr Pro Ser Lys Pro Ala Ser Ala Asp Gly His Arg
                            40
gge eee age geg gee tte gee eee geg gee gee gag eee aag etg tte
                                                                     192
Gly Pro Ser Ala Ala Phe Ala Pro Ala Ala Ala Glu Pro Lys Leu Phe
    50
                        55
```

					tcg Ser 70	_		_		_	_		 		240
_	_	-			gtg Val								_		288
		_			gac Asp										336
	_			_	gag Glu										384
					tac Tyr										432
		_	_		gag Glu 150										480
					ctc Leu									•	528
	_	_	-		acc Thr										576
/ -		_		_	aag Lys				 _						624
					ggc Gly										672
					gtg Val 230										720
					acc Thr										768
					gcc Ala										816
	_	_	_		cag Gln		_		 			_			864

	rp 1											ctg Leu 300					912
Me	_				_		_	_		_	_	gtc Val	_	_	_	_	960
												gtt Val					1008
				~	_			_	_	_		agt Ser	_	_	_		1056
												cct Pro					1104
	et A	_	_	_		_			_			gtg Val 380	_		_		1152
ΤΣ												ctg Leu					1200
			-			_	_			_	_	cgg Arg			_	_	1248
	_			_					_			ccc Pro				_	1296
-		Pro	_	_	_				_			atc Ile	_	_	_		1344
	cp S					_	_					aca Thr 460	_				1392
	co I											gac Asp					1440
												gag Glu					1488
												gag Glu					1536

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gag tac ctg cag gcc ttc ctg gag gac tac ttc acg tcc acc gag ccc
                                                                    1584
Glu Tyr Leu Gln Ala Phe Leu Glu Asp Tyr Phe Thr Ser Thr Glu Pro
        515
                            520
cag tac tag cccggggaga acctctag
                                                                    1611
Gln Tyr
    530
<210> 4
<211> 530
<212> PRT
<213> Homo sapiens
<220>
<221> MISC FEATURE
<222> (1)..(530)
<223> amino acid sequence of the mutant c-Src polypeptide encoded
       by the mutant c-Src coding region
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Ser Leu Glu Pro Ala Glu Asn Val His Gly Ala Gly Gly Ala Phe
Pro Ala Ser Gln Thr Pro Ser Lys Pro Ala Ser Ala Asp Gly His Arg
Gly Pro Ser Ala Ala Phe Ala Pro Ala Ala Ala Glu Pro Lys Leu Phe
    50
My Gly Phe Asn Ser Ser Asp Thr Val Thr Ser Pro Gln Arg Ala Gly
Pro Leu Ala Gly Gly Val Thr Thr Phe Val Ala Leu Tyr Asp Tyr Glu
                                    90
Ser Arg Thr Glu Thr Asp Leu Ser Phe Lys Lys Gly Glu Arg Leu Gln
            100
Ile Val Asn Asn Thr Glu Gly Asp Trp Trp Leu Ala His Ser Leu Ser
Thr Gly Gln Thr Gly Tyr Ile Pro Ser Asn Tyr Val Ala Pro Ser Asp
                        135
Ser Ile Gln Ala Glu Glu Trp Tyr Phe Gly Lys Ile Thr Arg Arg Glu
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Ser Glu Arg Leu Leu Asn Ala Glu Asn Pro Arg Gly Thr Phe Leu

170

165

Val Arg Glu Ser Glu Thr Thr Lys Gly Ala Tyr Cys Leu Ser Val Ser 180 185 190

Asp Phe Asp Asn Ala Lys Gly Leu Asn Val Lys His Tyr Lys Ile Arg 195 200 205

Lys Leu Asp Ser Gly Gly Phe Tyr Ile Thr Ser Arg Thr Gln Phe Asn 210 215 220

Ser Leu Gln Gln Leu Val Ala Tyr Tyr Ser Lys His Ala Asp Gly Leu 225 230 235 240

Cys His Arg Leu Thr Thr Val Cys Pro Thr Ser Lys Pro Gln Thr Gln
245 250 255

Gly Leu Ala Lys Asp Ala Trp Glu Ile Pro Arg Glu Ser Leu Arg Leu 260 265 270

Glu Val Lys Leu Gly Gln Gly Cys Phe Gly Glu Val Trp Met Gly Thr 275 280 285

Trp Asn Gly Thr Thr Arg Val Ala Ile Lys Thr Leu Lys Pro Gly Thr 290 295 300

Met Ser Pro Glu Ala Phe Leu Gln Glu Ala Gln Val Met Lys Lys Leu 305 310 315 320

Arg His Glu Lys Leu Val Gln Leu Tyr Ala Val Val Ser Glu Glu Pro 325 330 335

Ile Tyr Ile Val Thr Glu Tyr Met Ser Lys Gly Ser Leu Leu Asp Phe 340 345 350

Leu Lys Gly Glu Thr Gly Lys Tyr Leu Arg Leu Pro Gln Leu Val Asp 355 360 365

Met Ala Ala Gln Ile Ala Ser Gly Met Ala Tyr Val Glu Arg Met Asn 370 380

Tyr Val His Arg Asp Leu Arg Ala Ala Asn Ile Leu Val Gly Glu Asn 385 390 395 400

Leu Val Cys Lys Val Ala Asp Phe Gly Leu Ala Arg Leu Ile Glu Asp 405 410 415

Asn Glu Tyr Thr Ala Arg Gln Gly Ala Lys Phe Pro Ile Lys Trp Thr 420 425 430

Ala Pro Glu Ala Ala Leu Tyr Gly Arg Phe Thr Ile Lys Ser Asp Val 435 440 445

Trp Ser Phe Gly Ile Leu Leu Thr Glu Leu Thr Thr Lys Gly Arg Val 450 455 460

Pro Tyr Pro Gly Met Val Asn Arg Glu Val Leu Asp Gln Val Glu Arg 465 470 475 480

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Gly Tyr Arg Met Pro Cys Pro Pro Glu Cys Pro Glu Ser Leu His Asp
                485
                                    490
Leu Met Cys Gln Cys Trp Arg Lys Glu Pro Glu Glu Arg Pro Thr Phe
Glu Tyr Leu Gln Ala Phe Leu Glu Asp Tyr Phe Thr Ser Thr Glu Pro
Gln Tyr
    530
<210> 5
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<212> DNA
<213> Artificial Sequence
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<223> 3' mutant allele specific primer
<220>
<221> misc_feature
<222> (15)..(15)
<223> n is defined as a 3-nitropyrrole residue
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tagaggttct ccccnggcta
∢210>
<211>
      20
<212>
      DNA
<213> Artificial Sequence
<220>
<223> 3' wild-type allele specific primer
<400> 6
                                                                      20
tagaggttct ccccgggctg
<210> 7
<211> 164
<212> DNA
<213> Artificial Sequence
<220>
<223> Antisense sequence complementary to 5' region of c-Src gene
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<400> 7
gccccgcagg tgcctactgc ctctcagtgt ctgacttcga caacgccaag ggcctcaacg 60
tgaagcacta caagatccgc aagctggaca gcggcggctt ctacatcacc tcccgcaccc 120
agttcaacag cctgcagcag ctggtggcct actactccag tgag 164